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Title: NaCT AS A TARGET FOR LIFESPAN EXPANSION AND WEIGHT REDUCTION

Applicant(s): Ganapathy et al. Serial No.: Unassigned

Express Mail No.: EV 201890140 US

Filed: Herewith

Docket: 275.00080101 Sheet 1 of 44

Figure 1

1/44

drIndyl sequence (tol., 2602)

(ORF: 258 - 1976)

TTCACCGTTTCCGAATCGGACGAACCGGGCGTGATTGCTCTCCTGCTGCTTTCGAGATCGGAGTCCCGATAAGGATA TAACTACAACCTAAAGAGGAATCCAAGCCTCCTCCTGCCGCTAGTTTCGAAAAGTAAATAGAGTACTTGTTATCAAC TGGGAAGCGGAGATACATAGCTCCGATATTCCTGTGAAAGCCAGACAAACGGATACCAACGAACAATCGCCATATCT ACACGCCACCGCCACTGGACATCAAAATGGAAATTGAAATTGGCGAACAACCCCAGCCTCCGGTGAAGTGCTCCAAC TTCTTCGCTAACCACTGGAAGGGATTGGTTGTGTTCCTGGTGCCGCTGCTATGTCTGCCTGTTATGCTGCTAAACGA AGGCGCCGAATTTCGGTGCATGTACCTCCTTTTGGTAATGGCCATATTTTGGGTTACGGAAGCCTTGCCTCTATG TGACGTCCATGATACCGATTGTGGCCTTCCCAATAATGGGTATAATGAGCTCGGATCAGACTTGCCGCTTGTACTTC AAGGATACGCTGGTGATGTTCATGGGCGGCATTATGGTCGCCCTGGCTGTGGAGTACTGTAATCTACACAAACGTCT TGCCTTGAGGGTAATCCAGATCGTGGGCTGCAGTCCCCGCAGATTACACTTTGGCCTCATCATGGTTACAATGTTTT TGAGCATGTGGATTTCGAACGCCGCCTGTACTGCCATGATGTGTCCGATTATCCAAGCCGTGCTGGAGGAGCTGCAG GCTCAGGGTGTCTGCAAAATCAACCATGAGCCTCAATACCAAATCGTTGGAGGCAACAAGAAAAACAACGAGGATGA GCCACCATACCCCACCAAGATCACTGTGCTACTATCTGGGCATTGCCTACGCCTCCTCGCTGGGTGGCTGTGGAA CCATCATCGGAACTGCCACCAATCTTACCTTCAAGGGCATCTACGAGGCTCGTTTCAAGAACTCCACCGAACAGATG GACTICCCCACCTICATGTTCTACTCGGTGCCATCCATGTTGGTCTACACCTTGCTGACATTCGTGTTCCTGCAATG GCACTTCATGGGTCTGTGGCGTCCCAAGAGCAAGGAGGCACAGGAAGTCCAGAGGGGGACGAGAGGGCGCCGATGTCG CCAAAAAGGTTATCGATCAGCGCTACAAGGATCTGGGTCCCATGTCCATTCACGAGATCCAAGTGATGATTCTGTTC ATTITTATGGTTGTGATGTACTTCACCCGCAAGCCCGGCATCTTTTTGGGATGGGCCGATTTGCTGAATTCCAAGGA CATTCGTAACTCTATGCCCACTATTTTTGTCGTCGTCATGTGCTTCATGCTGCCCGCCAATTATGCTTTCCTACGCT ACTGCACCAGACGCGGTGGTCCAGTGCCCACGGGTCCCACTCCATCGCTGATCACCTGGAAGTTCATCCAGACCAAG GTGCCATGGGGTCTGGTGTTCCTGCTTGGCGGTGGCTTCGCTTTGGCCGAAGGCAGCAGCAGCAGGCGGCATGGCCAA GCTGATTGGCAATGCTCTGATTGGATTGAAGGTTCTGCCCAACTCTGTCCTCTTACTGGTGGTCATCCTGGTGGCTG TGTTCCTGACCGCCTTCAGCTCCAATGTGGCGATTGCCAACATTATTATTCCCGTTCTGGCCGAGATGTCCCTGGCC ATTGAGATCCATCCTGTACCTGATCCTGCCCGCTGGCTTGGCCTGCAGTATGGCCTTCCACCTGCCGGTTAGTAC TCCGCCCAACGCTTTGGTTGCTGGCTATGCCAACATTAGGACGAAGGACATGGCCATTGCTGGAATCGGTCCGACCA TCATTACCATCATCACCCTGTTTGTTTTCTGCCAAACCTGGGGCCTGGTCGTCTATCCGAACCTTAACTCGTTCCCC AACATACCCGTCACAGCGATAAAGTTGAGGAAAATTTAGGGGAATTTTAAACGAAAAGTGCCTTTGCTGACAGCGAAA **ANTGTGAAAAATATTTAACTATGTATACTTGCATTTCAGAGTTGCGAAAAGTTTTGATACAAAAGCATTACCTACTG** TTTAGAAAAATGTGTTAAAAAAAAAACGTATCGCAATATACTGTTAATCAGGAATTGAACACCTGGTCTACGCACTC AGCTAAATATTTAAATACAAATTAATGTTACTTAATTGTTGCATTTAGCATAAAAATGGAAAAGATTTGGAAAAGTT AGAACAGTTTGTTCAATGGCAGCCCTGGCCTGCTAATATTTTAAATAACTAGACTGAGAGAACTTACATATTCATAC TAGCGGTAGGCTAAGCTTAAATGATACTGTGTACATTTTCAGATGATTTATGTTTTATATAGTTTGTAAAAAAATATT

SEQ ID No:1

Peptide seq (total 572)

MEIEIGEQPQPPVKCSNFFANHWKGLVVFLVPLLCLPVMLLNEGAEFRCMYLLLVMAIFWVTEALPLYVT SMIPIVAFPIMGIMSSDQTCRLYFKDTLVMFMGGIMVALAVEYCNLHKRLALRVIQIVGCSPRRLHFGLI MVTMFLSMWISNAACTAMMCPIIQAVLEELQAQGVCKINHEPQYQIVGGNKKNNEDEPPYPTKITLCYYL GIAYASSLGGCGTIIGTATNLTFKGIYEARFKNSTEQMDFPTFMFYSVPSMLVYTLLTFVFLQWHFMGLW RPKSKEAQEVQRGREGADVAKKVIDQRYKDLGPMSIHEIQVMILFIFMVVMYFTRKPGIFLGWADLLNSK DIRNSMPTIFVVVMCFMLPANYAFLRYCTRRGGPVPTGPTPSLITWKFIQTKVPWGLVFLLGGGFALAEG SKQSGMAKLIGNALIGLKVLPNSVLLLVVILVAVFLTAFSSNVAIANIIIPVLAEMSLAIEIHPLYLILP AGLACSMAFHLPVSTPPNALVAGYANIRTKDMAIAGIGPTIITIITLFVFCQTWGLVVYPNLNSFPEWAQIYAAAALGNKTH

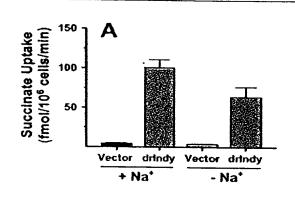
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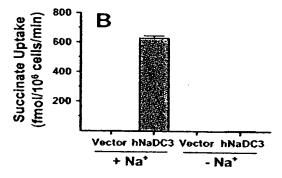
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Figure 2



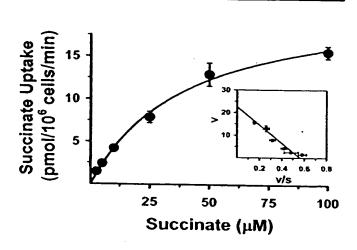


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Figure 3



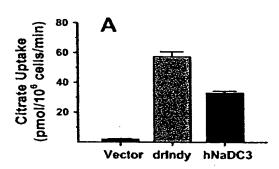
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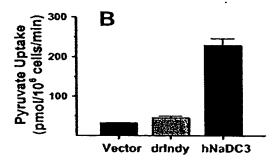
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Figure 4





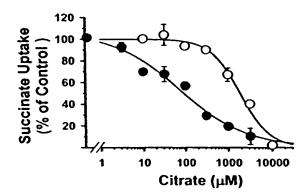
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Figure 5



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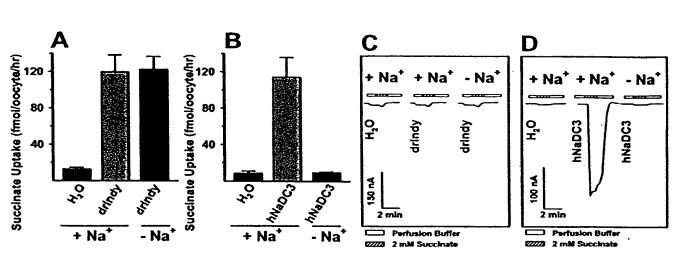


Figure 6

Title: NaCT AS A TARGET FOR LIFESPAN EXPANSION AND WEIGHT REDUCTION

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Figure 7

Rat INDY seq. (3191 nt + 63 nt polyA)

CCAGTCTGTCTCCCTTTCACGCGATGGCTTCGGCGAAGACTTATGTGACCAAGTTCAAGTCCTTCGTGATTTTGTTCTTCGCCCC GATCCTGCTGCTTCCACTCATCATTCTGGTACCTGACAAGTTTGCCAGGTGTGCCTATGTTATAATCCTCATGGCCATCTACTGG TATGTGTCCAATACATGACGGACACCAACATGCTGTTCCTGGGCAGTCTCATTGTGGCCACGGCTGTGGAACGTTGGGAAC7TCA TARGAGGATTGCTCTGAGAATGCTACTCTTTGTGGGGACCAAGCCTTCACGGCTGATGCTGGGCTTCATGTTCGTCACAGCCTTC CTGTCCATGTGGATCAGCAATACTGCCACCACGGCCATGATGATACCCATTGTGGAGGCCATGCTGGAGCAAATGGTAGCCAACA TGAAGACCCCAGTGTGCAGAAGCAGGAGGATGAAGAAACAAAGAATATGTACAAGGCTATGAACCTATGTGTGCTACGCAGCC **AGCATCGGGGGTACAGCACCTTGACCGGGACGGGACCCAACGTGGTGCTGCTGGGCAATGCAGGAATTGTTTCCTGACAGTA AAGACGTCATGAACTTTGCATCTTGGTTTGCCATTTGCCCTCCCAAACATGCTTTTGATGCTGGTGATGGCCTGGCTGTGGCTCCT** GTGTTTTTP.CATGAGACCCAATTTAAAAAAAACTTGCATCTGCTGGGGGAAGAAGAAGAAGGACACGGAGAAGATTCCCTCCAAG GTGCTGTATGAGGAGTACAGGAAGCTGGGGCCCTTGAGCTACGCTGAATGCAACGTGCTCTTTTGCTTCGGCCTGCTCATCATCC CACAGTGGCCATCTTTGTGGCCATTTTGCTTTTCATCGTACCCTCACAAAAGCCCCAAGTTCAATTTCAGCCGCCAGACTGAGGAA GAAAGGAAAACTCCCTTCTACCCCCCGCCACTGCTGAATTGGAAAGTCACCCAAGAGAAAGTGCCCTGGGGCATTGTGCTGCTCC TGGGGGGAGGATTTGCTATGGCCAAAGGATGTGAGACTTCAGGGCTCTCTGAGTGGATGGCGAGACAGATGGCGCCCTTGAGCTC **AGTGAGACCTGCTATTATTACCTTGATCTTGTCCTGTATTGTTGCAATGACCAGAGTGCACGAGTAACGTGGCCACTACTACC** CACTTGCCTTCATGTTGCCTGTGGCCACCCCACCTAACGCCATCGTGTTTGCCTACGGACACCCTCAAAGTTATTGACATGGTAAA ABATTCCCTGACTGGGCAAATTTGACACATATTAACACTTAGGAGAACCACAAGAGCACAGGCTTGTCCCCCAACCCTTTCGAGG ACTGCGAACCTTCTGGCACACCTTGCACAGAGCACTGGTGCTCATACCCCAGTGTGACCCCAATGATGTCAACACCCCAAGAAGAT CCTTTGAGAGGTCGTGAGGCCCATCTTCCCTAGGACCCTTCCATCTCACCTGGGCAGAAACAGAGGGACTGGGGCTCAAGTCCT GTACCACGTGGCTTTGAAAGACTTCTGACTCCATGCTGGGCTCTGGTTCTCACATGCCTGTTCCCACGGTCTCCACATGGGGATC AGATGACQAGAGACAGCCCCTGTGCCTCTTCTGGATGTTCCCAGATCACCATCTCTATCACACAGAAACTTCCTCTCTCACA **ACAGAACTCTGATCTTGAACACTTTCCACTGCCAGAGTTAGAGTGGAAATCACGGCCCCCTGAAGACTTTGACTCTACATGGTGC** CATCTCCAACCACTGGGCAACCTGAGGTGCTAACATTGAGGCCTTCCTGCTCACCCTTGGCTGACCTGTTCCCTACTTGCCTTAT . CTCTTATTAGTTAACAGTTLGAGGCCCCTTCCCAGCTCCCAGLGAGACTTCATCAACTCCTAGATGCTCCTGGCTGAGGCTCCC CAGAAAGTCCCTGTTTGATTGTTCTGGGGATGCTCATCCTGCCAGCCTGGAGCAGCTGGGTCATACATCAGGGATGGACAATGGT TRAGGGCTGGGCCACCACAGTTGGTCCTGTTGGACTCTTGAGCTCCTCCAGAAGGCCTTTTCCTTGCTGTGTGCACTGTGGGC GGT&G>GTCCATGGGA&CCCATGGGACCACTCACATGAAAGGGAGAGAGAGAGAAGAAGCTCTCCCTTGTCCTTCAGGGATGCT CTCTTCCTTGCTTAATTTGCTCTGAAAAGAAGCATGAGTGGGGAGATAAGATCCGTGGATGTCATTTTATTTTTCCAGGCAGAAAG AAGGCTTGCTCTTGTCAAATACTCTGTCTGCTATGGAAAGTTCCAGTGTGCTGACTGGTCTGTGATTTCGTGCCTTGTGAAGGAG AAAAAAAAAAAAAAAAAAAAAAA SEQ ID NO:3

(572 aa)

MASAKTYVTKFKSFVILFFAPILLIPLIILVPDKFARCAYVIILMAIYWCTDVIPVAITSLLPVLLFPLLKVLDSKQ VCVQYMTDTNMLFLGSLIVATAVERWELHKRIALRMLLFVGTKPSRLMLGFMFVTAFLSMWISNTATTAMMIPIVEA MLEQMVATHVAVDASQRTMELLDKNKASELPGSQVVFEDPSVQKQEDEETKNMYKAMNLCVCYAASIGGTATLTGTG PNVVLLGQMQELFPDSKDVMNFASWFAFALPNMLLMLVMAWLWLLCFYMRPNLKKTCICCGRKKKDTEKIASKVLYE EYRKLGPLSYAECNVLFCFGLLIILWFSRDPGFMPGWLSIAWIEGNTKHVTDATVAIFVAILLFIVPSQKPKFNFSR QTEEERKTPFYPPPLLNWKVTQEKVPWGIVLLLGGGFAMAKGCETSGLSEWMARQMEPLSSVRPAIITLILSCIVAM TTECTSNVATTTLFLPIFASMARSIGIHPLYVMIPCTLSASLAFMLPVATPPNAIVFAYGHLKVICMVKTGLVMNIL GIASVFLSVNTWGRAVFNLDKFPDWANLTHINT

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Figure 8

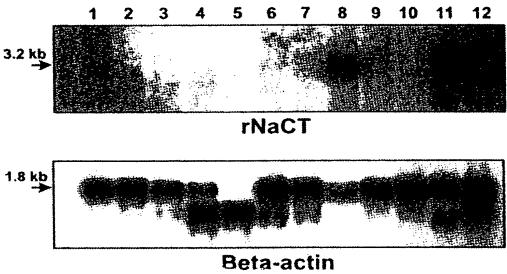
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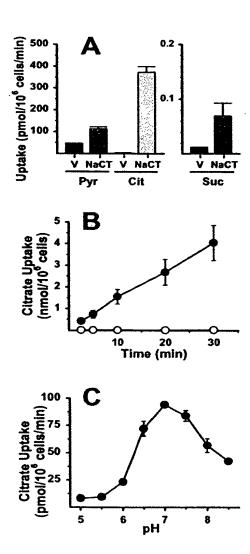
Figure 9



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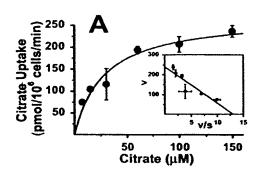
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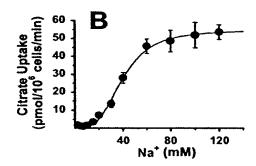
Figure 10

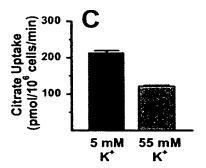


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Figure 11

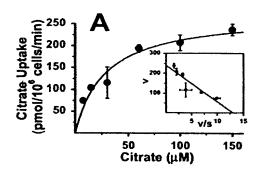


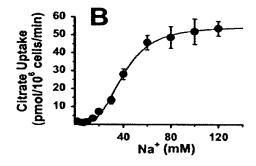


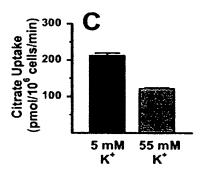


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Figure 12

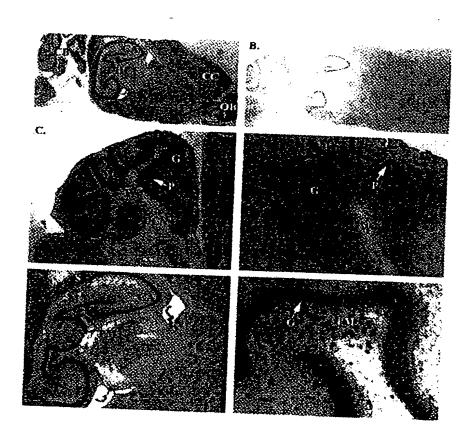






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Figure 13



Title: NaCT AS A TARGET FOR LIFESPAN EXPANSION AND WEIGHT REDUCTION

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Figure 14

Human NaCT sequence

(3207 nt + 41 nt polyA) ORF: 13-1719 (total 1707 nt)

CCCTCCCGCGCG

ATGGCC?CGGCGCTGAGCTATGTCTCCAAGTTCAAGTCCTTCGTGATCTTGTTCGTCACCCCGCTCCTGCTGCTGCCACTCC TCATTCTGATGCCCGCCAAGTTTGTCAGGTGTGCCTACGTCATCATCCTCATGGCCCATTTACTGGTGCACAGAAGTCATCCC ATGAAGGACACCAACATGCTGTTCCTGGGCGCCTCATCGTGGCGCTGTGGAGCCTGGAACCTGCACAAGAGAACATC CCCTGCGCACGCTCCTCTGGGTGGGGGCCAAGCCTGCACGGCTGATGCTGGGCTTCATGGGCGTCACAGCCCTCCTGTCCAT CTGGATCAGTAACACGGCAACCACGGCCATGATGGTGCCCCATCGTGGAGGCCATATTGCAGCAGATGGAAGCCACAAGCCCAC GCCACCGAGGCCGGCCTGGAGCTGGTGGACAAGGCCAAGGCCAAGGAGCTGAGGGAGTCAAGTGATTTTTGAAGGCCCCCA CTCTGGGGCAGCAGGAAGACCAAGAGCGGAAGAGGTTGTGTAAGGCCATGACCCTGTGCATCTGCTACGCGCCCAGCATCGG GGCACCGCCACCCTGACCGGGACGGGGACCCAACGTGGTGCTCCTGGGCCAGATGAACGAGTTGTTTCCTGACAGCAAGGAC CICGIGAACTITGCTICC1GG1TIGCATTTGCCTTTCCCAACATGCTGGTGATGCTGCTGTTCGCCTGGCTGTGCTCCAGT GCTGCAGGAGGAGTACCGGAAGCTGGGGGCCCTTGTCCTTCGCGGAGATCAACGTGCTGATCTGCTTCTTCCTGCTGGTCATC ATGCCACTGTGGCCATCTTTGTGGCTACCCTGCTATTCATTGTGCCTTCACAGAAGCCCAAGTTTAACTTCCGCAGCCAGAC GTGCTGCTACTAGGGGGGGGGATTTGCTCTGGGCTAAAGGATCCGAGGCCTCGGGGGCTGTCCGTGTGGATGGGGAAGCAGATGG AGCCCTTGCACGCAGTGCCCCCGGCAGCCATCACCTTGATCTTGTCCTTGCTCGTTGCCGTGTTCACTGAGTGCACAAGCAA CGTGGCCACCACCACCTTGTTCCTGCCCATCTTTGCCTCCATGTCTCCATCGGCCTCAATCCGCTGTACATCATGCTG CCCTGTACCCTGAGTGCCTCCTTTGCCTTCATGTGCCTGTGCCACCCTCCAAATGCCATCGTGTTCACCTATGGGCACC TCAAGGTTGCTGACATGGTGAAAACAGGAGTCATAATGAACATAATTGGAGTCTTCTGTGTGTTTTTTGGCTGTCAACACCTG GGGACGGGCCATATTTGACTTGGATCATTTCCCTGACTGGGCTAATGTGACACATATTGAGACTTAG GAAGAGCCACAAGACCACACACACAGCCCTTACCCTCCAGGACTACCGAACCTTCTGGCACACCTTGTACAGAGTTTTGG GGTTCACACCCCAAAATGACCCAACGATGTCCACACACCACCACCCAAAACCCAGCCAATGGGCCACCTCTTCCTCCAAGCCCAGA TGCAGAGATGGACATGGGCAGCTGGAGGGTAGGCTCAGAAATGAAGGGAACCCCTCAGTGGGCTGCTGGACCCATCTTTCCC AAGCCTTGCCATTATCTCTGTGAGGGAGGCCAGGTAGCCGAGGGATCAGGATGCAGGCTGCTGTACCCGCTCTGCCTCAAGC ATCCCCCACACAGGGCTCTGGTTTTCACTCGCTTCGTCCTAGATAGTTTAAATGGGAATCAGATCCCCTGGTTGAGAGCTAA GACAACCACCTACCAGTGCCCATGTCCCTTCCAGCTCACCTTGAGCAGCCTCAGATCATCTCTGTCACTCTGGAAGGGACAC CCCAGCCAGGGACGGAATGCCTGGTCTTGAGCAACCTCCCACTGCTGGAGTGGGAATCAGAGCCTCCTGAAGCCTC AGTGCACCACAGGCCAACCTACGCCCTTCATCACTTGGTTCTGTTTTAATCGACTGGCCCCCTGTCCCACCTCTCCAGTGAG TICTCCCAGGCAGGTCATCTTTTCTGGGAGATGCTTCCCCTGCCATCCCAAATAGCTAGGATCACACTCCAAGTATGGGCA ACTGGCTATGCCACTICAGAGICTITCATGCCAGCGTTTGAGCTCCTCTGGGTAAAATCITCCCTTTGTTGACTGGCCTTCA CAGCCATGGCTGGTGACAACAGAGGATCGTTGAGATTGAGCAGCGCTTGGTGATCTCTCAGCAAACAACCCCTGCCCGTGGG CCAATCTACTTGAAGTTACTCGGACAAAGACCCCAAAGTGGGGCAACAACTCCAGAGAGGCTGTGGGAATCTTCAGAAGCCC CCCTGTAAGAGACAGACATGAGAGACAAGCATCTTCTTTCCCCCGCAAGTCCATTTTATTTCCTTCTTGTGCTGCTGGAA OKCOCO TO TO TO THE TOTAL PROPERTY OF THE TO ALGGACCGAGTATGTGTGGGTTCCTTGGGTGGGACGATTCCTGACCACACTGTCCAGCTCTTGCTCTCATTAPATGCTCTG

Amino acid sequence (568 aa)

MASALSYVSKFKSFVILFVTPLLLLPLVILMPAKFVRCAYVIILMAIYHCTEVIPLAVTSLMPVLLFPLFQILDSRQVCVQY PROTINHEFEGGLI VAVAVERHNEHKRI ALRTLEHVGAKPAREHLGFMGVTALLSMHI SNTATTAMMV PI VEA I LQQMEATSA ATEAGLELVDKGKAKELPGSQVIFEGPTLGQQEDQERKRLCKAHTLCICYAASIGGTATLTGTGPNVVLLGQMNELFPDSKD Lynfashfafafphmlymllfahlhlqfvymrfnfkkshgcgleskknekaalkylqeeyrklgplsfaeinvlicffllvi LXFSRDPGFM:PGWLTVAWVEGETXYVSDATVAIFVATLLFIVPSQXPXFNFRSQTEEERKTPFYPPPLLDHKVTQEKVPWGI VLLLGGGFALAXGSEASGLSVHHGKQMEPLHAVPPAAITLILSLLVAVFTECTSHVATTTLFLP1FASHSRSIGLNPLYIHL PCTLSASFAFMLPVATPPNAIVFTYGHLKVADHVKTGVIHNI IGVFCVFLAVNTHGRAIFDLDEFPDHANVTHIET

SEQ ID NO:6

SEO ID No:2

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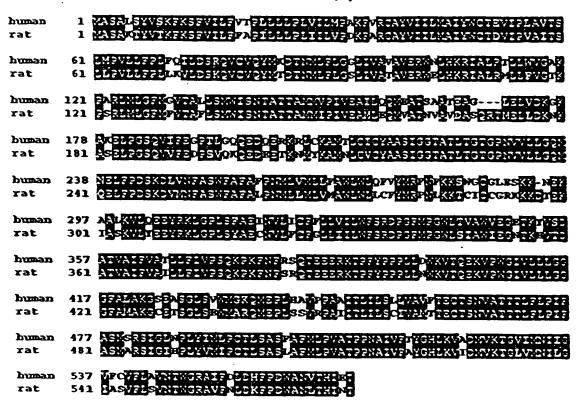
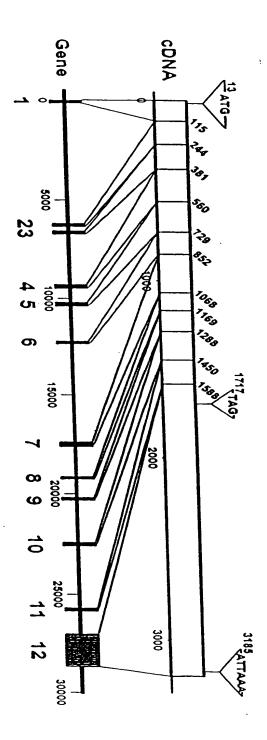


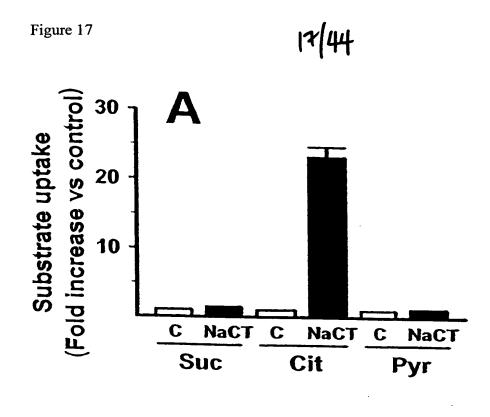
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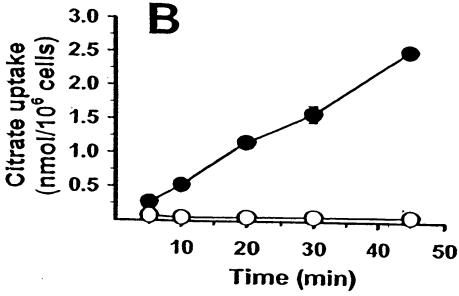
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Figure 16

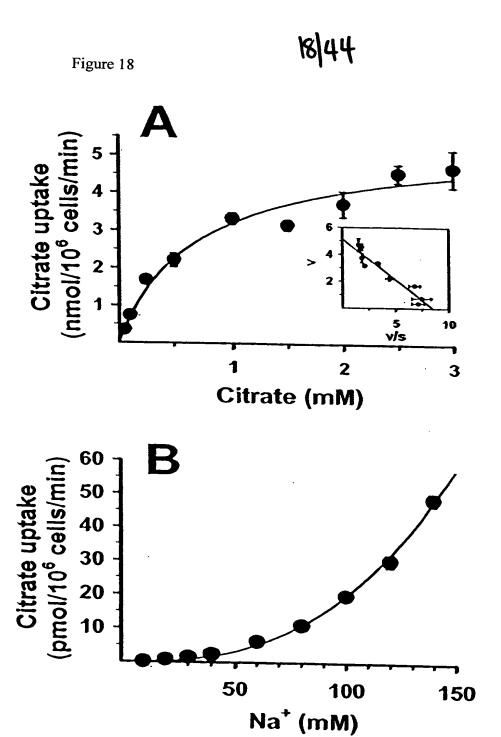


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Figure 19

ceNaCT cDNA & Protein Sequences

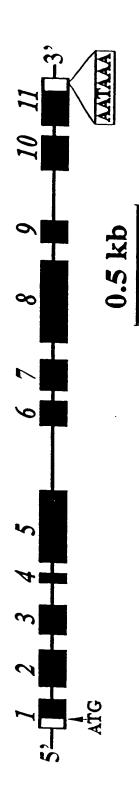
1	CTTTGAGTGTTACCACTATCAGACGAAGGTATGAAGCCTAGCCCCCAGCGTACGTTAATAAAAAAGCTTCTGGTACTTCTTGGACCACTTGTTGCAGTAC	
101	M R P S P Q R T L I R R L L V L L G P L V A V P CTCTACTCTTTTTTGGACCGGAGTACAGATOCTTTTCTCAATAATATTTTTATCAACCTATTGGATGGAGAAGCATTTCCCATTGGTGTCACTTCTCT	100
201	L L F F G P E Y R C L F S I I F L S T Y W I G E A F P I G V T S L CTTTCCATTGGCACTTTATCCAATTCTTCAGATTGTTCCATCTAAACAAATTAGTCCAGTTTATTTTAAAGATTCGATAGTTCTATTTATGTGCACATTA	200
301	PPLALYPILQIVPSKQISPVYPKDSIVLPMCTLAGCATGGCAATGCCGCTGGAATCGCGCTGAATTATTAACAAAAGTTGGAGCAAAGCAACCAGTAATGCTGCTGGTT	300
401	S M A M A V E A T G L H R R I A L R L L T R V G A R Q P V M L L G P TCATGTGCATCACGAGTTTCATATCATTTTTCGTTTCTGACACACGCATGCACACCCCTGTGGCACTCCTGATGAGTATGTCTGA	400
501	M C I T S F I S F F V S D T A C T A L M C P T A V A L L M S M S D TOCAGTTCAACATTTGAAAGAAGATCACAGGAAGCCCAAAGCCCACAGATGATGCTACTGTTGCTCAAAAAATGAGATGATATGACTCCTCAG	500
601	A V Q H L K E D H R K P K P P P D D A T V A E K M R I D D M T P Q GATGCTGGTTTCTGTAAAGCATTAATTTTGGCATGTCCCCACGCATCGTTGATCGGTGGGACTGCTATTATCACTTCGACTGGGCCGAATTTAGTTTTTC D A G P C K A L L L A C A H A C C C A C C C C	600
	D A G F C K A L I L A C A H A S L I G G T A I I T S T G P N L V F R GAGAAAATATTCACAAGAGATACCCCGAGGGACAAGTGACAATGACATGCTACCAATGGATGCTTTCGCGATACCACCAATGTTTGTCTACCTTCTTGC	700
	ENIHERYPEGQVTMTYLQWMVFAIPPMFVYLLA ATCTTATATTATTCTGGTGTGTTATTCATGGGCCCCTCGACTTTTGCTCGCTGGTTTGAAAGGCCCTTCCAAAGAAGAAGCTCATTTGAAAAAATTAATT	800
	GAAAAGAATATTCAGACAATGTATGAGGAGTTTTCGGGGGGGG	900
	E K N I Q T M Y E D L G D V S W G E K S V F V F P I L L I G S W I S CTGGTGATCCGGGATTCACACCGGATCGGGAGACTTTTTACCACACAGAAACTTCATATCAGACAGTGTTTCTGGAGTCTTGATTTCTGTATTTTATT	1000
	R D P G F T P G W G D L L P H R N F I S D S V S G V L I S C I L F TGITTGGCCCAAAGATCCATTGATCCTATTGATCCAATGCTCCCATTCTTAAATGGACCGACATGAAAAGCAAGTTTTCCTGGTCGTGCACACTTTTA	1100
1201	V W P K D P F D P I D P H A P I L K W T D H K S K F S W S C T L L ATCOGTOCTOGGTATCCTATTTCAGAAGGAGTTGATAAATCGGGATTATCTAGATTGATT	1200
1301	TICCATTGCAATTAACTGTGACTACAATTATTGTGATAATGACAAGAGTTTGCAAGTAATGTGTCCACCGGAAGCATTTTCATTCCAATTTCTTTTGGAGT	1300
	TGCTGAATCAATGGGAGTTCATCCTTTATATCTTGCTCTTCCAACTACTGTTCATTTGCTTTTATGCTACCAATTTCAACACCTCCAAATGCA	1400
1501	GTTGTATACGACACTAAAGTGATTTCAATGGTGGAAATGATGGTGGTTTTCTACTCAATATTGCGTGCATACTTATCACATCTCTCAACATGAACA	1500
1601	V V Y D T K V I S M V E M I V C G P L L N I A C I L I T S L N M N T CATGGACATATTTTATTTTTCATTGAATATTTTCCCCGAAAATATTGTAATATCTTCAGAAAACTCTTCATATCCAGTTTGCTAATTTTTGTACAAAAT	1600
1701	H T Y P I P S L N I P P E N I V I S S E N S S Y P V C • GTGTATTGTCCGAATGAAACGTGTATTTATTAAAAAAAAA	1700

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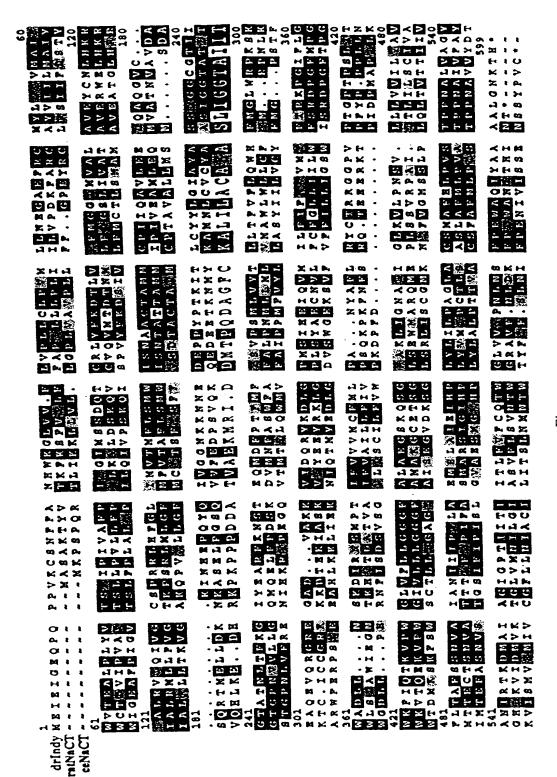
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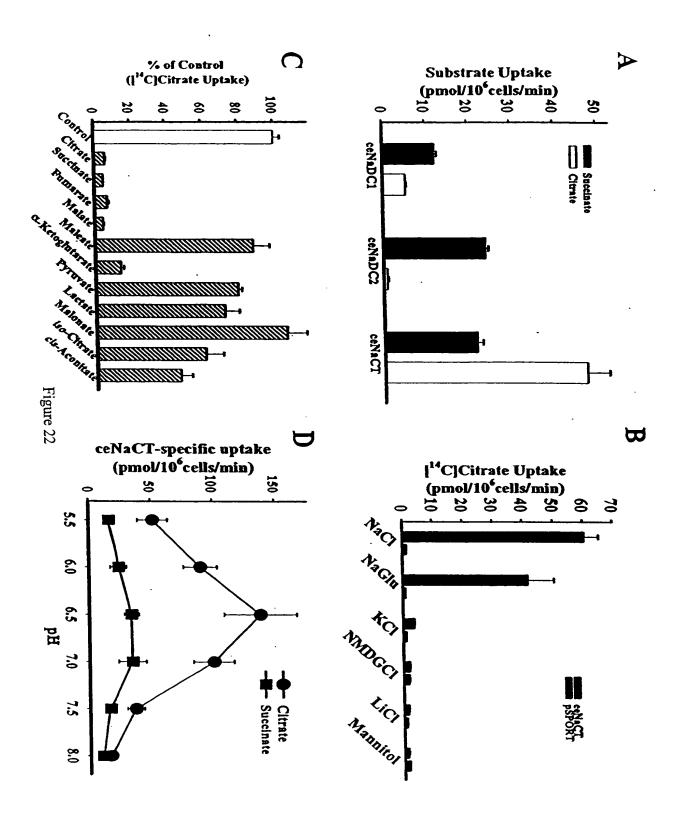
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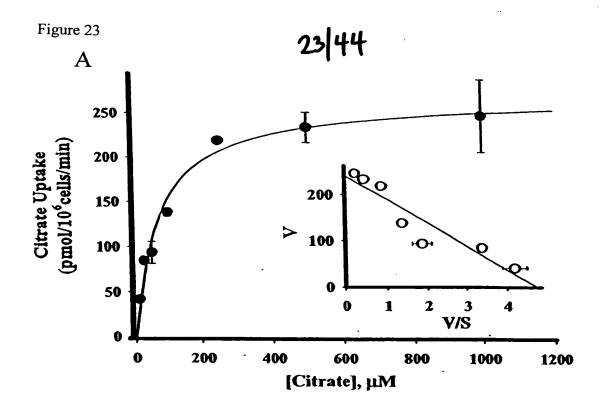
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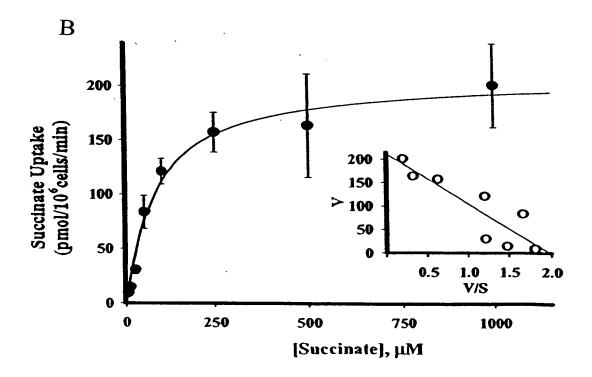
Figure 2

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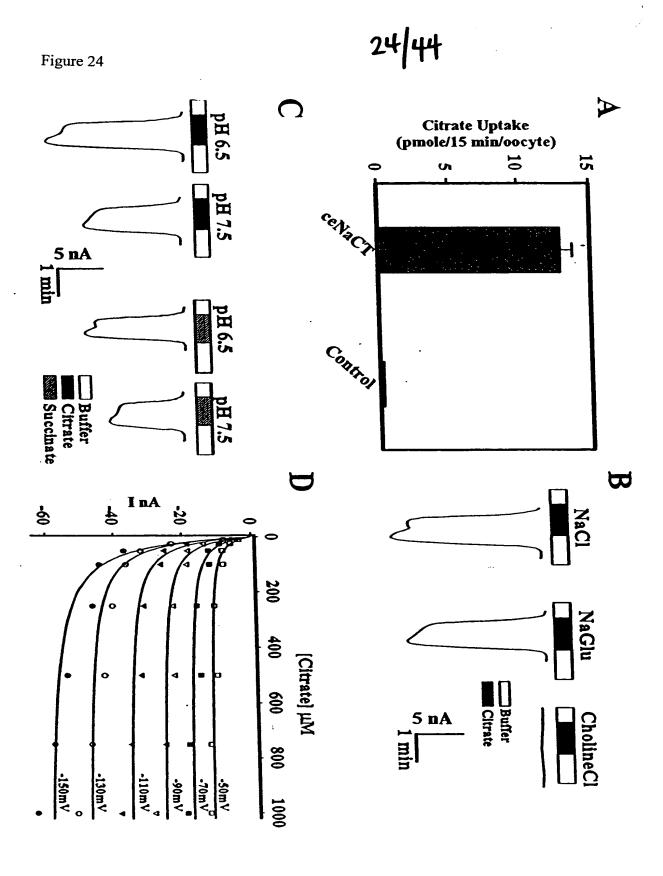


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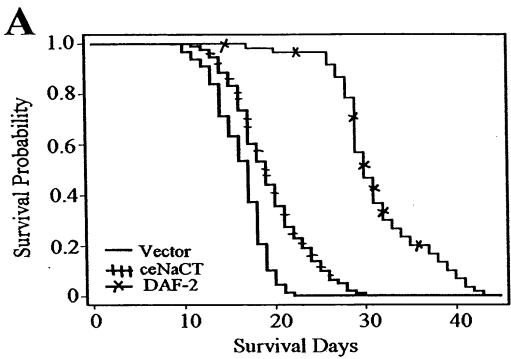
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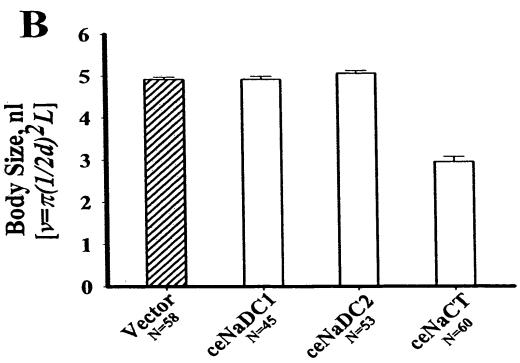


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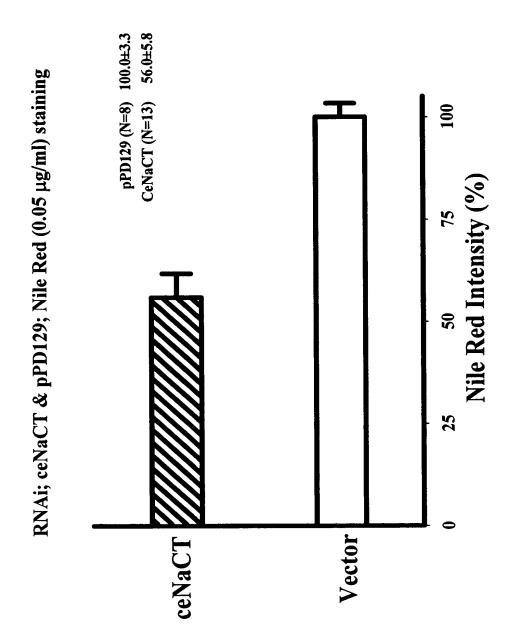


Figure 26

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Figure 27

SEQ ID NO:9

Mouse NaCT sequence

cDNA sequence (16 nt + 1719 nt)

GTCTCCCTTTCACGCG

ATGGATTCGGCGAAGACTTGTGTGACCAAGTTCAAGTCCTTTGCGATTTTGCTCTTCACCCCGATCCT GATGCTTCCACTCGTCATTCTGATACCTGACAAGTTTGCCAGGTGTGCCTATGTTATAGTCATTATGG CTTTTGAAGGTTCTGGACTCCAAGCAGGTATGTATCCAATACATGAAGGACACCAACATGCTGTTCCT GGCAGTCTCATTGTGGCTGTGGCTGTGGAACGCTGGAAACTTCATAAGAGGGTTGCCCTGAGAATGC-TGCTCTTTGTGGGGACCAAGCCCTCACGGCTGATGCTGGGCTTTATGTTTGTCACGGCCTTCCTGTCC ATGTGGATCAGCAATACTGCCGCCACAGCCATG ATGATACCCATTGTGGAGGCCATGCTGCAGCAAAT GATAGCCGCCAATACAGCTGTGGAGGCCCAGCCTGGGGCACACTGGAGCTGCTGGACAAGAACAAGACCA GCGAGTTGCCAGGAAGCCAGGTGGTATTTGAAGACCCCAATGTGCAGGAGCAGGAAGACGAAGAACA GACCGGGACGGACCCAACGTGGTGCTCCTGGGCCAGATGCAGGAATTGTTTCCTGACAGTAAAGATG TCCTGAACTATGCATCTTGGTTTGGATTTGCCTTCCCCAACATGGTGATGATGCTGGTGCTGGCCTGG GAGGGACACCGAGAAGATTGCCTACAAAGTGCTGAACGAGGAGTACCAGAAGCTGGGGTCCTTGAGCT ACCCTGAATGCAACGTGCTCTTTTGCTTCACCCTACTTGTCATCCTGTGGTTCTCCCGAGACCCCGGC TTCATGCCTGGCTGTCATTCGCCTGGGTCGAGGGAAACACCGTTCATATCACAGATGCCACAGT GGCCATCTTTGTGGCCATTTTGCTTTTCATCATACCTTCACAAAAGCCCCAAGTTCAACTTCAGCAGCC AGACTGAGGAAGAAAGGAAAACTCCGTTCTACCCCCCAGCACTGCTGGATTGG AAAGTCGCCCAAGAG AGGGCTCTCGAAGTGGATGGCAGCAGATGGAACCCTTGAGATTAGTGAAACCTGCTGTCATTACCT TGATCTTGTCCTGTCTTGCAATGACCACAGAGTGCACAAGTAACGTGGCCACTACCACCCTGTTC CTGCCTATCTTTGCCTCCATGGCTCGTTCCATTGGTA TCCATCCTCTGTATGTCATGATTCCCTGTAC CATGAGTGCTTCACTTGCCTTCATGTTGCCTGTGGCCACCCCACCGAATGCCATCGTGTTTGCCTACG GACACCTCAGAGTTGTTGACATGATGAAAACAGGATTGATAATGAACTTCGTTGGAATCCTATCTGTG TTTCTGTCAGTCAACACCTGGGGTCGGGCTATGTTTAACTTGGATAACTTCCCCGACTGGGCAAATTC **AACAÁGTGTTAACACTTAG**

Protein sequence (572 nt)

SEG ID NO:10

MDSAKTCVTKFKSPAILLFTPILMLPLVILIPDKFARCAYVIVIMAVYWCTDVIPVAVTSLLPVLLFPLLKVLDSKQV
CIQYMKDTNMLFLGSLIVAVAVERWKLHKRVALRMLLFVGTKPSRLMLGFMFVTAPLSMWISNTAATAMMIPIVEAML
QQMIAANTAVEASLGTLELLDKNKTSBLPGSQVVFEDPNVQEQEDEBTKNMYKAMHLCVCYSASIGGTATIGTGPNV
VLLGQMQELFPDSKDVLNYASWFGFAPPNMVMMLVLAWLWLQCLYMRHNLKKTCICCGBKKRDTEKIAYKVLNBBYQK
LGSLSYPECNVLFCFTLLVILWFSRDPGFMPGWLSFAWVEGNTVHITDATVAIFVAILLFIIPSQKPKFNFSSQTEBB
RKTPFYPPALLDWKVAQEKVPWDIVLLLGGGFAMAKGCBTSGLSKWMAAQMEPLRLVKPAVITLILSCLVAMTTECTS
NVATTTLFLPIFASMARSIGIHPLYVMIPCTMSASLAFMLPVATPPNAIVFAYGHLRVVDMMKTGLIMNFVGILSVFL
SVNTWGRAMFNLDNFPDWANSTSVNT

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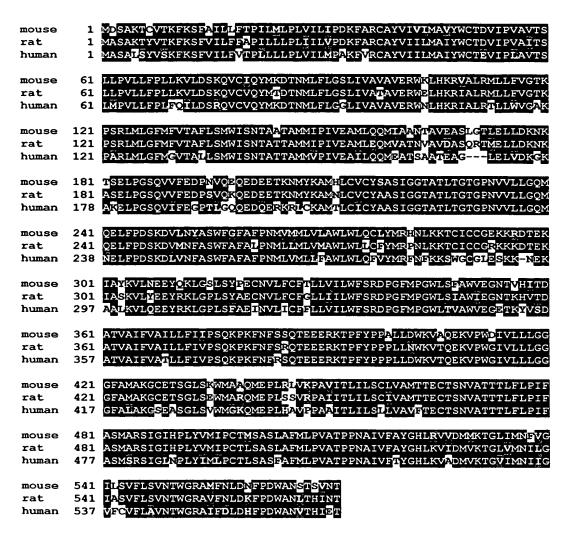
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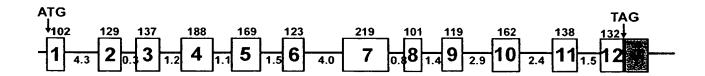
Figure 28

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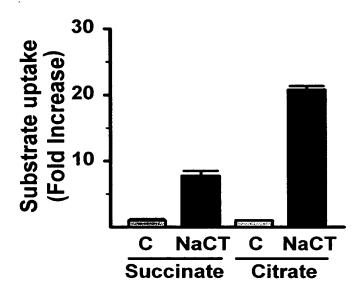


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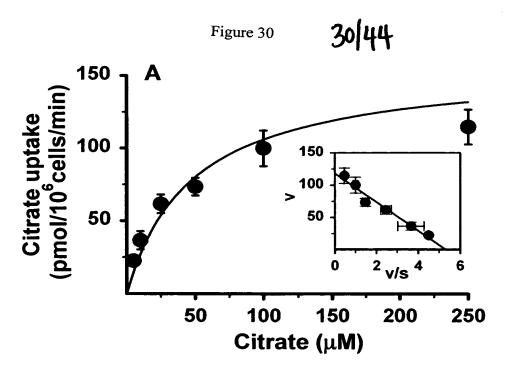


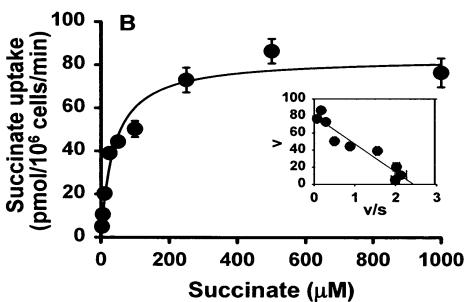
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Figure 29



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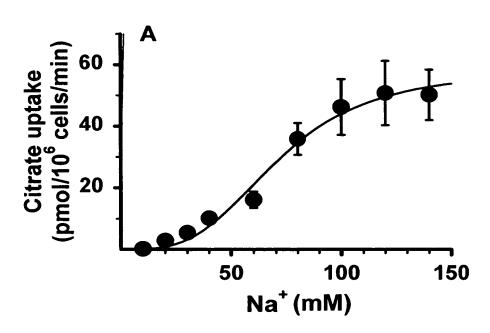


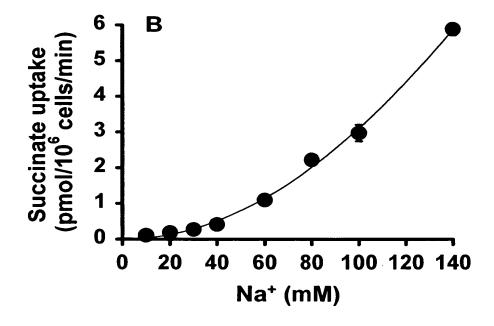


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Figure 31



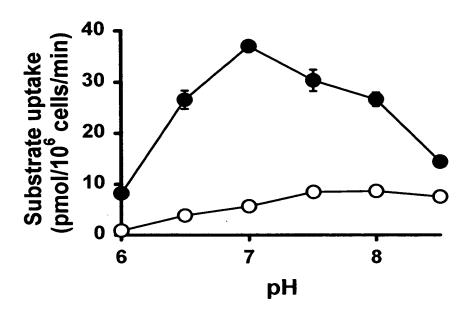


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Figure 32

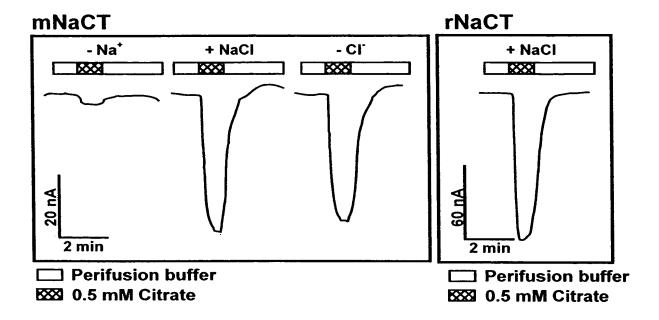


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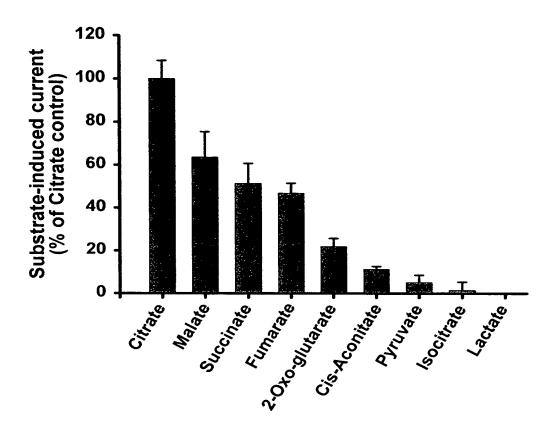
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Figure 33

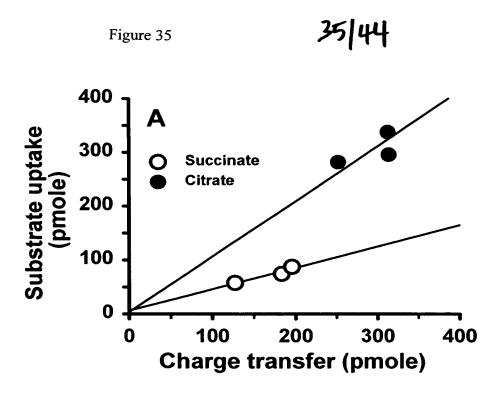


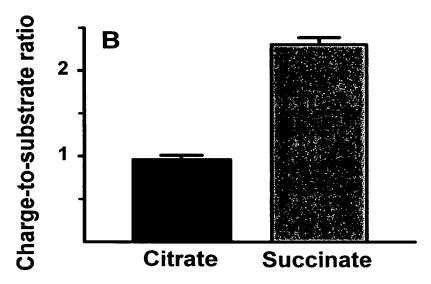
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Figure 34



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Figure 36

Zebra Fish NaCT full length cDNA (1#) (1-2536 + 15 bp)

ORF: from 76 - 1824 (length = 1749) (SEQ IO NO 11)

AGAGATCAGCGCACAGAAGTTTTGCGCAGTTTCTCACCGTTTGGACATTTCATTGTAAAGTTATCCAAAGCCGAAA TO GCTTCACGTGCACTCAAACTAGTATGGAAGATGAAAAATACATTGATTCTTTTTTGCACTCCATTTCTTCT GCTTCCTTTGCCACTTGTCATTGGATCAAAGGAGGCTGGATGTGCATATGTTGTGGTACTGATGGCAGTTTACTGG TGTACAGAGGTGCTGCCGCTGGCTGTCACTGCTCCTCCCGCCGCTGTGCTCTTTCCCCTCTTCAGAATCATGGAGT CCCAAGACGTATGTATGCAGTACCTTAAGGACACTAACATGCTGTTTCTGGGTGGCCTGATGGTGGCCGTGGCTGT CGAACACTGGAATCTGCACAAGCGGATCGCCCTGCGGGTGCTCCTTGTTGGGGGGTTCGACCAGCTCTGTTAATG TTGGGCTTCATGGGTGTAACAGCTTTCCTCTCCATGTGGATCAGTAACACGGCCACAACAGCCATGATGGTGCCCA TCGTTCAGGCAGTTCTCGAGCAGCTCAACAACACAGCACAACAAGAACAAAGCTCCATACCTGAGACCGAGGAAAA GAGCACTGAGAAACAGCCTGAGAGCCCGGGTGAGGAAAAAGTGGTACTGAATGGCGACAACTTCTCAATGGAGTCA GACCCTGAAGAACATTCACGAGAAGCAGAGGAAAGGCTGAAGATGTCTAAAGGCCTGACCCTGTGCGTGTTATG CCGCCAGCATCGGCGGCACACCCACACTCACAGGCACTGGACCAAACCTCGTTCTTATGGGACAGATGAGCCAACT GTTCCCGGACAACCCTGACATCATTAACTTTGCGTCATGGTTTGGATTTGCCTTTCCAAACATGATCATCATGCTC ACGCTGGCCTGGCTGTGCTACAGATCGTGTTTCTGGGAATAAACTTTAAAAAGACATGGGGCTGTGGGACGGTGA AGACGGAGAAGGAGATCGCGGCCTATAATGTGATTAAAGAGGAGCACCGCAGTCTCGGCCCTATGACCTTTGGGGA GCTGAGTGTCCTTGCCCTCTTCATCCTCCTGGTGGTGCTTTGGTTCACTCGTGATCCAGGCTTCGTGGACGGCTGG GCGACACGCTTCTTCAATGCTGACAAAGAGTTTGTGACAGATGCCACGGTTGCAGTGTTTGTGGCTGCTCT TTGTCTTTCCCTCTAAACCACCACGATTGTGCTTCTGGAGAACAGAGAGTTTCGACACAGTGCCCCAGCAAGAAAG TGGCCCGACTCCAGCTTTGCTGACATGGAAAGTGACACAGAAGAAGATGCCATGGAGTATTATACTGCTGCTGGGA GGAGGCTTTGCCCTGGCTAAGGGCAGTGAGATCTCAGGATTGTCCAAGTGGCTTGGAGATCAGATGTCTCCTCTTC AAAGCATTCCTCCATGGGCAATAGCTATTGTCATATGTTTAATGATCGCAACCTTCACTGAATGCACCAGTAATGT GGCCACAGCTACATTATTTCTGCCTATACTGGCATCTATGTCTCAGTCTATAGGTGTGAATCCTCTGTATGTTATG GTGCCCTGTACCCTCAGTGCATCTTTTGCCTTCATGCTCCCTGTGGCAACTCCTCCAAACGCCATCGTCTTCTCAT ACGGATACCTCAAAGTCTCTGACATGGCCAAGACTGGGATCGTCATGAACATCATCGGCATCCTCTCCATCACCTT AGCCATTAACAGCTGGGGCAGAGCCATCTTCAGTTTAGACACGTTCCCCAGCTGGGCAAACACTACTGATGTC GAGACACAGAGAGCCGGACTGCCCCACTCACCACTTGTGAACTTCAGATTGTTTCCAGTTCTCATGTGAACAGAGA AAACCGATTATGACCACTGTTTAGTCATTTCACACATTCATGTCTATCTTTTACAAAACCGTGGTGTTCACTACAG TCTGTGGACATTGTAACACCAATATTTGAATGATTCATAAAAGACAAAAATAGATTTTACAAATCATGATTTTTCT GTAGCATTACAGCATACTGTGAGCACTGAGCATATATTTGGACCATTGGTTGTTCGCTTTTGGCTTGTGAAAGAGC ATTTGGACGCAGAAACTCAACATCAGTCATATTGTCCCCTTGTGTCCAGACTCAGAGCCAGTGGCCGGTTTCACTG GAGATCAATACTGTACTTTGACGGTTCAAATCACTTCATGTTCATATGCGTAATTTAAAGCTGCTTTAAACACAGC TACACAAGAGCACATGCAGAAAAAGCTGAAAAGGCTGAAATTGTGTAAATATCATCTAGATTTTTAAGCACAAAAGT ATCATGTACATTTCAAAATTAATTCAAACTCTATTTTTAATGTACATTTATTAAAATTATGTATTTTGTTCATATT TACT ΑΑΑΑΑΑΑΑΑΑΑΑ

Amino Acid Sequence (581 aa) (SEQ IÛ NO: 12)

MASRALKLVWKMKNTLILFCTPFLLLPLPLVIGSKEAGCAYVVVLMAVYWCTEVLPLAVTALLPAVLFPLFRIMES QDVCMQYLKDTNMLFLGGLMVAVAVEHWNLHKRIALRVLLLVGVRPALLMLGFMGVTAFLSMWISNTATTAMMVPI VQAVLEQLNNTAQQEQSSIPETEEKSTEKQPESPGEEKVVLNGDNFSMESDPEEHSREAEERLKMSKGLTLCVCYA ASIGGTATLTGTGPNLVLMGQMSQLFPDNPDIINFASWFGFAFPNMIIMLTLAWLWLQIVFLGINFKKTWGCGTVK TEKEIAAYNVIKEEHRSLGPMTFGELSVLALFILLVVLWFTRDPGFVDGWATRFFNADKEFVTDATVAVFVAALLF VFPSKPPRLCFWRTESFDTVPQQESGPTPALLTWKVTQKKMPWSIILLLGGGFALAKGSEISGLSKWLGDQMSPLQ SIPPWAIAIVICLMIATFTECTSNVATATLFLPILASMSQSIGVNPLYVMVPCTLSASFAFMLPVATPPNAIVFSY GYLKVSDMAKTGIVMNIIGILSITLAINSWGRAIFSLDTFPSWANTTDV

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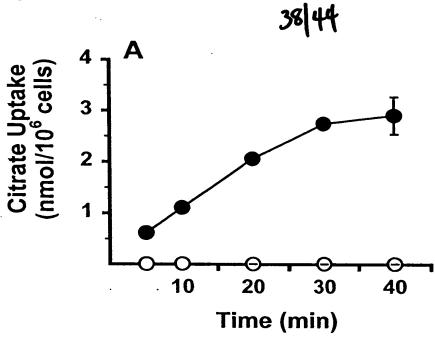
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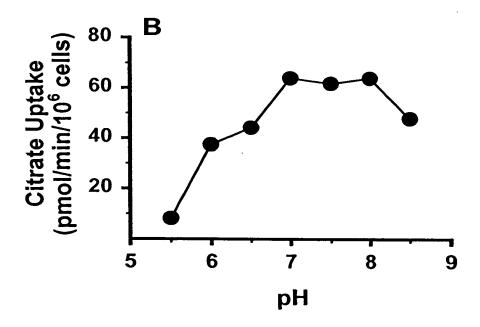
Figure 37

fish	1 MASRALKLVWKMKNTLILFCTPFLLLPLBLVIGSKBAGCAYVVVLMAVYWCTEVLPLAVI
mouse	1 -MDSAKTCVTKFKSFAILLFTPILMLPLVILIPDKFARCAYVIVIMAVYWCTDVIPVAVI
rat	1 -MASAKTYVTKFKSFVILFFAPILLDPLIILVPDKFARCAYVIILMAIYWCTDVIPVAIT
human	1 -MASALSYVSKFKSFVILFVTPLLLDPLVILMFAKFVRCAYVIILMAIYWCTEVIPLAVT
fish mouse rat human	61 ALLFAVLFPLFRIMESODVCMQYLKDTNMLFLGCLMVAVAVEHWNLHKRIALRVLLLVGV 60 SLLPVLLFPLLKVLDSKQVCIQYMKDTNMLFLGSLIVAVAVERWEHKRIALRVLLFVGT 60 SLLPVLLFPLLKVLDSKQVCVQYMTDTNMLFLGSLIVATAVERWEHKRIALRNLLFVGT 60 SLMPVLLFPLFQLLDSRQVCVQYMKDTNMLFLGGLIVAVAVERWNLHKRIALRTLLWVGA
fish	121 RFALLMLGFMCVTAFLSMWISNTATTAMMVPIVQAVLEQINNTAQQEQSSIPETEEKSTE
mouse	120 KESRLMLGFMFVTAFLSMWISNTAATAMMIPIVEAMLOQMIAANTAVEASLGTLELLDKN
rat	120 KESRLMLGFMFVTAFLSMWISNTATTAMMIPIVEAMLEQMVATNVAVDASQRIMELLDKN
human	120 KFARLMLGFMCVTAILSMWISNTATTAMMVPIVEAILQQMEATSAETEAGLELVDKG
fish	181 KOPESPGEBKVVLNGDNFSMESDFEEHSREABERLKMSKGLTLCVCYAASIGGTATLTGT
mouse	180 KISELPGSQVVFEDPNVQBQEDETKNMYKAMHLCVCYSASIGGTATLTGT
rat	180 KASELPGSQVVFEDPSVQKQEDEETKNMYKAMNLCVCYAASIGGTATLTGT
human	177 KAKELPGSQVIFEGPTLGQQEDQERKRICKAMTLCICYAASIGGTATLTGT
fish	241 GPNLVLMGQMSOLFPDNPDIINFASWFGFAFPNMIIMLILAWLWLCIVFLGINFKKTWGC
mouse	231 GPNVVLLGQMQBLFPDSKDVLNYASWFGFAFPNMVMMLVLAWLWLQCLYMRHNIKKTCIC
rat	231 GPNVVLLGQMQBLFPDSKDVMNFASWFAFALPNMLLMLVMAWLMLICFYMRPNIKKTCIC
human	228 GPNVVLLGQMNBLFPDSKDLVNFASWFAFAFPNMLVMLLFAWLMLCFVYMRFMFKKSWGC
fish	301 G-TVKTEKEIAAYNVIKEEHRSLGPMTFGELSVIALFILLVVLWFTRDPGFVDGWATR-F
mouse	291 CGEKKRDTEFIAYKVLNEEYOKLGSLSYBECNVLFCFTLLVILWFSRDPGFMPGWLSBAX
rat	291 CCRKKKDTEKIASKVLYEEYRKLGPLSYAECNVLFCFCLLIILWFSRDPGFMPGWLSIAW
human	288 GLESKK-NEKAALKVLQEEYRKLGPLSFAEINVLICFFLLVILWFSRDPGFMPGWLTVAW
fish mouse rat human	359 FNADKEFVTDATVAVEVAALLEVEPSKEPRLCEWRTESFDTVPQQESGPTEALLTWKVTC 351 VEGNTVHITDATVAIFVAILLEIJPSQKPKENESSQTEEER?TPFYPEALLDWKVAC 351 JEGNTKHVTDATVAIFVAILLEIVPSQKPKENESRQTEEER?TPFYPEPLLMWKVTC 347 VEGETFYVSDATVAIFVATLLEIVPSQKPKENERSQTEEER?TPFYPEPLLDWKVTC
fish mouse rat human	419 KMMPWSIILLLGGGFALAKGSEISGLSKWLGDQWSPLOSIPFWAIAIVICLMIATFTECT 408 EKVPWDIVLLLGGGFAMAKGCETSGLSKWMAAQMEPLRIVKPAVITLILSCLVAMTTECT 408 EKVPWGIVLLLGGGFAMAKGCETSGLSEWMARQMEPLSSVRPAIITLILSCIVAMTTECT 404 EKVPWGIVLLLGGGFALAKGSEASGLSVWMGKQMEPLHAVPPAAITLILSLLVAVFTECT
fish mouse rat human	479 SNVATATLFLPILASMSQSIGVNPLYVMVPCTLSASFAFMLPVATPPNAIVFSYGYLKVS 468 SNVATTTLFLPIFASMARSIGIHPLYVMIPCTMSASLAFMLPVATPPNAIVFAYGHLRVV 468 SNVATTTLFLPIFASMARSIGIHPLYVMIPCTLSASLAFMLPVATPPNAIVFAYGHLKVI 464 SNVATTTLFLPIFASMSRSIGINPLYIMLPCTLSASFAFMLPVATPPNAIVFTYGHLKVA
fish mouse rat human	539 DMAKTGIVMNIIGILSI <mark>T</mark> AINSWGRAIFSLETFFSWANTTDV 528 DMMKTGLIMNEVGILSVFLSVNTWGRAMFNLENFPDWANSTSVNT 528 DMVKTGLVMNIIGIASVFLSVNTWGRAVFNLDKFPDWANLTHINT 524 DMVKTGVIMNIIGVFCVFLAVNTWGRAIFDLDHFPDWANVTHIEI

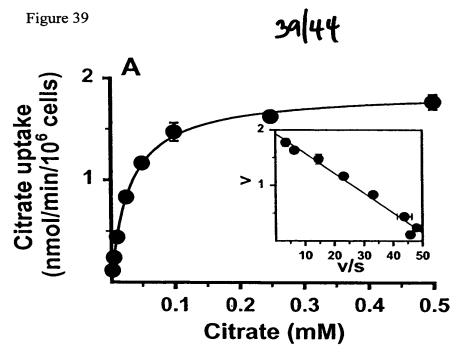
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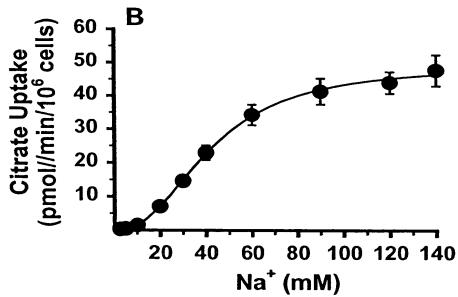




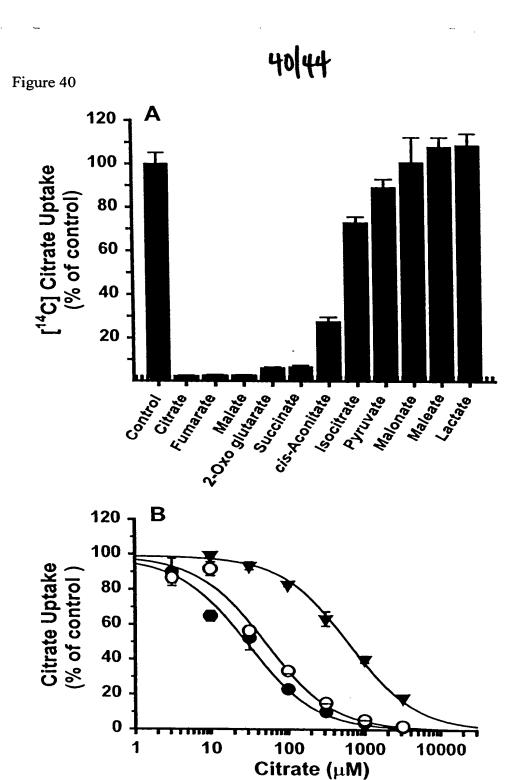


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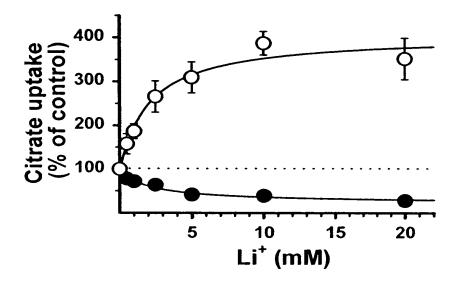


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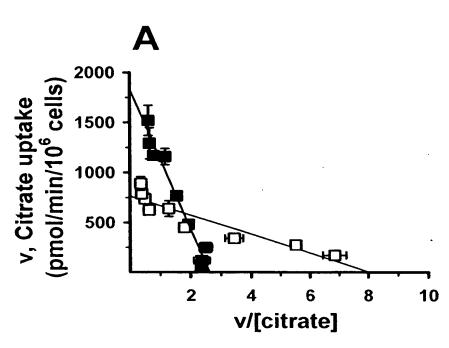
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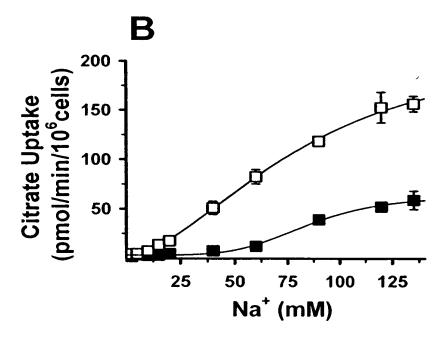


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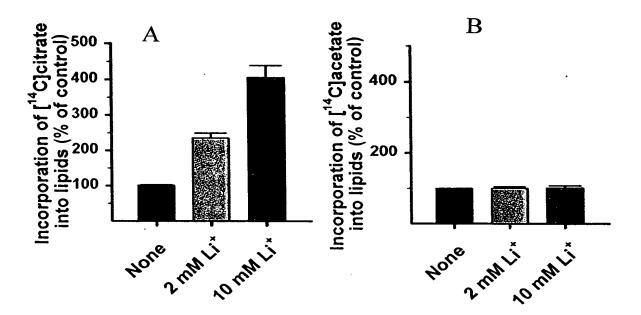


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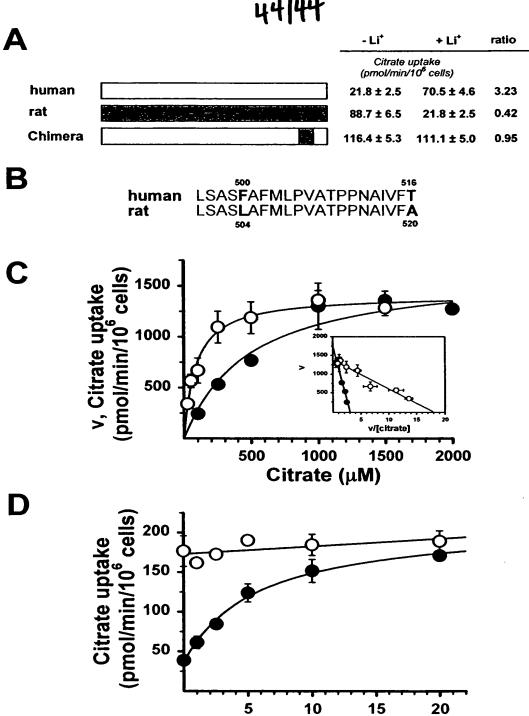
Figure 43



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Figure 44

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Li⁺ (mM)